IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicant(s): Gleave, et al. Application No.: 09/913,325 Filed: 8/10/2001 Title: TRPM-2 Antisense Therapy Attorney Docket No.: UBC.P-020 Customer No.: 57381	Group Art Unit: 1635 Examiner: Tracy Vivlemore Confirmation No: 8469

Commissioner for Patents
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SUPPLEMENT TO AMENDMENT ACCOMPANYING RCE

Sir:

Supplemental to the Amendment Accompanying RCE previously filed March 21, 2007, attached is the information regarding PC-3 cells referred to in the amendment.

Respectfully,

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Note: Performing your original search, *+"pc-3" "androgen receptor"*, in PubMed will retrieve *176* citations.

1: J Steroid Biochem Mol Biol. 2003 Apr;84(5):493-502.

Secretion of endogenous kallikreins 2 and 3 by androgen receptor-transfected PC-3 prostate cancer cells.

Kollara A, **Diamandis EP**, **Brown TJ**.

Samuel Lunenfeld Research Institute, Mt. Sinai Hospital, Suite 876, 600 University Avenue, Toronto, Ont., Canada M5G 1X5.

Androgen independent PC-3 cells lack androgen receptor (AR) expression and do not produce kallikrein 2 (hK2) or 3 (prostate-specific antigen, PSA). In this paper, we examined the ability of androgens to stimulate PSA and hK2 production in AR transfected PC-3 cells (PC-3(AR)) and compared this to LNCaP cells. PSA and hK2 were measured in the culture medium and cell lysates using an ELISA-based immunofluorometric assay. Only androgens were able to induce PSA and hK2 secretion in PC-3(AR) cells in a dose- and time-dependent manner depending on the level of AR present. The level of androgen-induced PSA and hK2 secretion in PC-3(AR) cells was approximately 1.5 and 0.9% that induced in LNCaP cells, respectively. Insulin-like growth factor-I (IGF-I), which has been shown to activate AR in the absence of ligand, did not activate PSA secretion in the absence of androgen, but further increased the dihydrotestosterone-induced PSA secretion in PC-3(AR) cells. The lack of PSA and hK2 production in parental PC-3 cells is thus a result of their lack of AR expression. PSA and/or hK2 production in PC-3(AR) cells can thus serve as an endogenous reporter system to investigate AR action or to screen putative endocrine disrupters.

PMID: 12767274 [PubMed - indexed for MEDLINE]

Related Links

Dissociation between androgen responsiveness for malignant growth vs. expression of prostate specific differentiation markers PSA, hK2, and PSMA in human prostate cancer models. [Prostate. 2003]

Androgen receptor activation in prostatic tumor cell lines by insulin-like growth factor-I, keratinocyte growth factor, and epidermal growth factor. [Cancer Res. 1994]

Interactive effects of triiodothyronine and androgens on prostate cell growth and gene expression. [Endocrinology. 1999]

Different proportions of various prostate-specific antigen (PSA) and human kallikrein 2 (hK2) forms are present in noninduced and androgen-induced LNCaP cells. [Prostate. 2000]

Tumor necrosis factor-alpha represses androgen sensitivity in the LNCaP prostate cancer cell line. [J Urol. 2000]

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